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## United States Patent [19]

### Oppermenn et al.

### Patent Number:

5,266,683

Date of Patent:

Nov. 30, 1993

#### [54] OSTEOGENIC PROTEINS

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[ \* ] Notice:

The portion of the term of this patent subsequent to Nov. 2, 2010 has been

disclaimed.

[21] Appl. No.: 841,646

[22] Filed:

Feb. 21, 1992

#### Reinted U.S. Application Data

Continuation-in-part of Ser. No. 827,052, Jan. 28, 1992, Pat. No. 5,250,302, Ser. No. 579,865, Oct. 7, 1990, Pat. No. 5,108,753, Ser. No. 621,849, Dec. 4, 1990, alandoned, Ser. No. 621,988, Dec. 4, 1990, abandoned, Ser. No. 810,560, Dec. 20, 1991, abandoned, Ser. No. 567,920, Aug. 20, 1990, abandoned, Ser. No. 600.024, Oct. 18, 1990, abandoned, Ser. No. 599,543, Oct. 18, 1990, shandoned, Ser. No. 616,374, Nov. 21, 1990, Pat. No. 5,162,114, and Ser. No. 483,913, Feb. 22, 1990, Pat. No. 5,171,374, said Ser. No. 827,052, is a division of Ser. No. 179,406, Apr. 8, 1988, Pat. No. 4,968,590, said Ser. No. 579,863, is a division of Ser. No. 179,406, Apr. 1, 1988, said Ser. No. 621,849, is a division of Ser. No. 232,630, Aug. 15, 1988, ubandoned, which is a continuation-in-part of Ser. No. 179,406, Aug. 15, 1988, said Ser. No. 621,988, is a division of Ser. No. 315,342. Feb. 23, 1989, Pat. No. 5,011,691, which is a continuation-in-part of Ser. No. 232,630, Feb. 23, 1989, said Ser. No. 810,560, is a continuation of Ser. No. 660,162, Feb. 22, 1991, abandoned, which is a continuation of Ser. No. 422,699, Oct. 17, 1989, abandoned, which is a continuation-in-part of Ser. No. 315,342, Oct. 17, 1989, said Ser. No. 569,920, is a continuation-in-part of Ser. No. 422,699, Oct. 17, 1989, and Ser. No. 483,913, Oct. 17, 1989, which is a continuation-in-part of Scr. No. 422,613, Oct. 17, 1989, Pat. No. 4,975,526, which is a continuation-in-part of Ser. No. 315,342, Oct. 17, 1989, said Ser. No. 600,024, is a continuation-in-part of Ser. No. 569,920, Oct. 17, 1989, said Ser. No. 599,543, is a continuation-in-part of Ser. No. 569,920, Oct. 17, 1989.

[51]	Int. Cl. <sup>3</sup> A-1K 37/02; C07K 5/C0; C07K 7/00; C07K 15/00
[52]	U.S. Cl 530/326; 530/327;
[58]	530/328; 530/350; 530/395; 530/840 Field of Search

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#### ABSTRACT

Disclosed are (1) osteogenic devices comprising a matrix containing substantially pure natural-sourced mammalian osteogenic protein; (2) DNA and amino acid sequences for novel polypeptide chains useful as subunits of dimeric octeogenic proteins; (3) vectors carrying sequences encoding these novel polypeptide chains and host cells transfected with these vectors; (4) methods of producing these polypeptide chains using recombinant DNA technology; (5) antibodies specific for these novel polypeptide chains; (6) osteogenic devices comprising these recombinantly produced proteins in association with an appropriate carrier matrix; and (7) methods of using the osteogenic devices to mimic the natural course of endochondral bone formation in mani-

58 Claims, 47 Drawing Sheets



